AMENDMENTS TO THE CLAIMS

Claims 1-29 (Cancelled)

30. (New) A method of reducing a compound of general structure III,

wherein X represents either hydrogen or OR₂,

and wherein R_1 and R_2 may be the same or different and represent hydrogen, or a hydroxy protecting group,

in an inert solvent with a chiral reducing agent or with a reducing agent in the presence of a chiral auxiliary,

3

to give a mixture of compounds of general structure IVa and IVb,

Application No.: NEW

Docket No.: 3893-0230PUS2

which is enriched with IVa, wherein X, R_1 , and R_2 are as defined above.

31. (New) A method for producing calcipotriol $\{(5Z, 7E, 22E, 24S)-24-\text{cyclopropyl-9,}10-\text{secochola-5,}7,10(19),22-\text{tetraene-}1\alpha-3\beta-24-\text{triol}\}$ or calcipotriol monohydrate comprising the steps of:

(a) reducing a compound of general structure III,

wherein X represents OR2,

and wherein R_1 and R_2 may be the same or different and represent hydrogen or a hydroxy protecting group,

in an inert solvent with a chiral reducing agent or with a reducing agent in the presence of a chiral auxiliary,

to give a mixture of compounds of general structure IVa and IVb,

which is enriched with IVa,

wherein X, R_1 and R_2 are as defined above;

(b) reacting the mixture of compounds of general structure IVa and IVb, which is enriched with IVa, in the presence of a base to give a mixture of compounds of general structure Va and Vb, which is enriched with Va,

wherein X, R₁ and R₂ are as defined above;

- (c) separating the compound of general structure Va from the mixture of compounds of general structure Va and Vb which is enriched with Va, wherein X, R₁ and R₂ are as defined above;
 (d) isomerising the compound of general structure Va to the compound of general structure VIa,
 - OH H VIa

wherein X, R₁ and R₂ are as defined above; and

(e) when R_1 and/or R_2 are not hydrogen, removing the hydroxy protecting group(s) R_1 and/or R_2 of the compound of general structure VIa to generate calcipotriol or calcipotriol monohydrate.

32. (New) A method for producing calcipotriol or calcipotriol monohydrate comprising steps (a) – (b) of claim 31 and further comprising the steps of:

(f) isomerising the mixture of compounds of general structure Va and Vb, wherein X, R₁ and R₂ are as defined in claim 2, which is enriched with Va, to a mixture of compounds of general structure VIa and VIb, which is enriched with VIa,

wherein X, R₁ and R₂ are as defined above;

(g) separating the compound of general structure VIa from the mixture of compounds of general structure VIa and VIb which is enriched with VIa, wherein X, R_1 and R_2 are as defined above; (h) when R_1 and/or R_2 are not hydrogen, removing the hydroxy protecting group(s) R_1 and/or R_2 of the compound of general structure VIa to generate calcipotriol or calcipotriol monohydrate.

33. (New) A method for producing calcipotriol $\{(5Z, 7E, 22E, 24S)-24-\text{cyclopropyl-9}, 10-\text{secochola-5}, 7, 10(19), 22-\text{tetraene-1}\alpha-3\beta-24-\text{triol}\}$ or calcipotriol monohydrate comprising the steps of:

(j) reducing a compound of general structure III,

wherein X represents hydrogen,

and wherein R_1 represents hydrogen or a hydroxy protecting group,

in an inert solvent with a chiral reducing agent or with a reducing agent in the presence of a chiral auxiliary,

to give a mixture of compounds of general structure IVa and IVb, which is enriched with IVa,

wherein X and R_1 are as defined above;

(k) reacting the mixture of compounds of general structure IVa and IVb, which is enriched with IVa, in the presence of a base to give a mixture of compounds of general structure Va and Vb, which is enriched with Va,

wherein X and R_1 are as defined above;

(1) separating the compound of general structure Va from the mixture of compounds of general structure Va and Vb which is enriched with Va, wherein X and R_1 are as defined above;

Docket No.: 3893-0230PUS2

Application No.: NEW

(m) hydroxylating the compound of general structure Va with a suitable hydroxylating agent, wherein X and R_1 are as defined above to give a compound of general structure Va, wherein X represents OR_2 and R_2 represents hydrogen, and wherein R_1 is as defined above;

(o) isomerising the compound of general structure Va to the compound of general structure VIa,

wherein X, R₁ and R₂ are as defined above; and

- (p) when R_1 is not hydrogen, removing the hydroxy protecting group R_1 of the compound of general structure VIa to generate calcipotriol or calcipotriol monohydrate.
- 34. (New) A method for producing calcipotriol or calcipotriol monohydrate comprising steps (j) (l) of claim 33 and further comprising the steps of:
- (q) protecting the C-24 hydroxy group of the compound of general structure Va,

wherein X represents hydrogen, and wherein R_1 represents hydrogen or a hydroxy protecting group, with a hydroxy protecting group;

- (r) hydroxylating the C-24 hydroxy protected compound of general structure Va with a suitable hydroxylating agent, wherein X and R_1 are as defined above to give a C-24 hydroxy protected compound of general structure Va, wherein X represents OR_2 and R_2 represents hydrogen, and wherein R_1 is as defined above;
- (s) removing the C-24 hydroxy protecting group of the compound of general structure Va;
- (t) isomerising the compound of general structure Va to the compound of general structure VIa,

wherein X, R₁ and R₂ are as defined above; and

- (u) when R_1 is not hydrogen, removing the hydroxy protecting group R_1 of the compound of general structure VIa to generate calcipotriol or calcipotriol monohydrate.
- 35. (New) The method according to claim 30, wherein the reducing step is with a reducing agent in the presence of a chiral auxiliary.
- 36. (New) The method according to claim 30, wherein the reducing agent is a borane derivative.
- 37. (New) The method according to claim 35, wherein the reducing agent is *N,N*-diethylaniline-borane, borane-tetrahydrofuran, or borane dimethylsulfide.
- 38. (New) The method according to claim 35, wherein the chiral auxiliary is a chiral 1,2-amino-alcohol.

Application No.: NEW

39. (New) The method according to claim 35, wherein the chiral auxiliary is a chiral *cis*-1-amino-2-indanol derivative.

Docket No.: 3893-0230PUS2

- 40. (New) The method according to claim 35, wherein the chiral auxiliary is (1S,2R)-(-)-cis-1-amino-2-indanol.
- 41. (New) The method according to claim 30, wherein the inert solvent is toluene, *tert*-butyl methyl ether, tetrahydrofuran, or mixtures thereof.
- 42. (New) The method according to claim 30, wherein the mixture of compounds of general structure IVa and IVb obtained by reducing a compound of general structure III has a molar ratio of IVa:IVb which is at least 56:44.
- 43 (New) The method according to claim 40, wherein the reducing step is carried out at a temperature between 10-20°C.

44. (New) A method for producing a compound of general structure III,

wherein X represents either hydrogen or OR2,

and wherein R_1 and R_2 may be the same or different and represent hydrogen, or a hydroxy protecting group,

by reacting a compound of general structure VII or VIII,

wherein R_1 and R_2 are as defined above, with sulphur dioxide.

45. (New) A method according to claim 30 or 44, wherein the compound of general structure III is the epimer of general structure IIIa

46. (New) A method according to claim 30 or 44, wherein the compound of general structure III is the epimer of general structure IIIb

47. (New) A method of reacting the mixture of compounds of general structure IVa and IVb,

wherein X represents either hydrogen or OR₂,

and wherein R_1 and R_2 may be the same or different and represent hydrogen, or a hydroxy protecting group,

which is enriched with IVa, in the presence of a base to give a mixture of compounds of general structure Va and Vb, which is enriched with Va,

wherein X, R_1 , and R_2 are as defined above.

- 48. (New) A method according to claim 30, 44, or 47, wherein X represents OR₂.
- 49. (New) A method according to claim 48, wherein R_1 and/or R_2 represent alkylsilyl.
- 50. (New) A method according to claim 48, wherein R₁ and/or R₂ represent tert-butyldimethylsilyl.

Docket No.: 3893-0230PUS2

Application No.: NEW

51. (New) A method for producing calcipotriol $\{(5Z, 7E, 22E, 24S)-24\text{-cyclopropyl-9,}10\text{-secochola-5,}7,10(19),22\text{-tetraene-}1\alpha\text{-}3\beta\text{-}24\text{-triol}\}\$ or calcipotriol monohydrate comprising the method of claim 30, 44, or 47.

52. (New) A compound of general structure IIIa or IIIb, or mixtures thereof,

wherein X represents either hydrogen or OR₂,

and wherein R_1 and R_2 may be the same or different and represent hydrogen, or a hydroxy protecting group.

53. A compound of general structure IVaa, IVab, IVba, IVbb, IVb, or mixtures thereof,

wherein X represents either hydrogen or OR₂,

and wherein R_1 and R_2 may be the same or different and represent hydrogen, or a hydroxy protecting group.

54. (New) A compound according to claim 52 or 53, wherein X represents OR_2 .

- 55. (New) A compound according to claim 54, wherein R_1 and R_2 represent alkylsilyl.
- 56. (New) A compound according to claim 54, wherein R_1 and R_2 represent *tert*-butyldimethylsilyl.
 - 57. (New) A compound according to claim 54, wherein R₁ and R₂ represent hydrogen.
- 58. (New) Use of a compound according to claim 52 or 53 as an intermediate in the manufacture of calcipotriol or calcipotriol monohydrate.